



427.045-DIV-2

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

In re Application of:

BIGG et al

Serial No.:

Filed: Concurrently Herewith

For: NEW...CONTAINING THEM

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: Examiner: S. Patel

:

: Group: 1624

:

600 Third Avenue

New York, NY 10016

**PRELIMINARY AMENDMENT**

Hon. Commissioner for Patents

P.O. Box 1450

Alexandria, VA 22313-1450

Sir:

Please amend this application as follows:

**Amendments to the Specification** begin on page 2 of this paper.

**Amendments to the Claims** are reflected in the listing of claims which begins on page 3  
of this paper.

**Remarks** begin on page 9 of this paper.

Mr PS. 07. check the  
Amendment date  
#11/03  
KFF  
4/26/04.

**New derivatives of N-(iminomethyl)amines, their preparation, their  
use as medicaments and the pharmaceutical compositions containing  
them**

--This application is a division of U.S. patent application Serial No. 09/787,466  
filed March 16, 2001, now U.S. Patent No. 6,482,822 which is a 371 of  
PCT/FR99/02251 filed September 22, 1999.--

A subject of the present invention is new derivatives of N-(iminomethyl)amines  
comprising the aminodiphenylamine, oxodiphenylamine, carbazole, phenazine,  
phenothiazine, phenoxazine or oxodiphenyl unit in their skeleton. These derivatives  
have an inhibitory activity on NO-synthase enzymes producing nitrogen monoxide NO  
and/or an activity which traps the reactive oxygen species (ROS). The invention relates  
to the derivatives corresponding to general formula (I) defined below, their preparation  
methods, the pharmaceutical preparations containing them and their use for therapeutic  
purposes, in particular their use as NO-synthase inhibitors and selective or non selective  
traps for reactive oxygen species.

Given the potential role of NO and the ROS's in physiopathology, the new derivatives  
described corresponding to general formula (I) may produce beneficial or favourable  
effects in the treatment of pathologies where these chemical species are involved. In  
particular:

- Proliferative and inflammatory diseases such as for example atherosclerosis,  
pulmonary hypertension, respiratory distress, glomerulonephritis, portal  
hypertension, psoriasis, arthrosis and rheumatoid arthritis, fibroses, angiogenesis,  
amyloidoses, inflammations of the gastro-intestinal system (ulcerous or non-ulcerous  
colitis, Crohn's disease), diarrhoea.
- Diseases affecting the pulmonary system and airways (asthma, sinusitis, rhinitis).
- Cardio-vascular and cerebro-vascular disorders including for example, migraine,  
arterial hypertension, septic shock, ischemic or hemorrhagic, cardiac or cerebral  
infarctions, ischemias and thromboses.